

Substitute for form 1449B/PTO <b>SUPPLEMENTAL INFORMATION IN CLOSURE STATEMENT IN AN APPLICATION LISTING OF REFERENCES</b> <b>June 4, 2009</b> (Use several sheets if necessary)	ATTORNEY DOCKET NO. 2345.2046-007		APPLICATION NO. 10/533,365	
	FIRST NAMED INVENTOR Inga Reynisdottir		371(c) DATE November 9, 2005	
	EXAMINER Juliet Caroline Switzer	CONFIRMATION NO. 4456	GROUP 1634	

Examiner's Initials	Ref. No.	OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)
	C5	Annex to European Search Report dated February 7, 2006 for EP Application No. 03783097.3-2405, filed 31 October 2003.
	C6	Supplemental Partial European Search Report dated October 30, 2007 for European Patent Application No. 03783097.3-1222, filed 31 October 2003.
	C7	Communication from EPO for Office Action for EP Application No. 03783097.3-1222, dated May 29, 2008.
	C8	Written Opinion of the International Preliminary Examining Authority for PCT/US03/34801, dated July 11, 2006.
	C9	International Preliminary Report on Patentability for PCT/US03/34801, dated September 11, 2006.
	C10	First Notification of Office Action for Chinese Application No. 2003801026117, dated February 15, 2008.
	C11	Communication from EPO for Office Action for EP Application No. 03783097.3-1222, dated March 9, 2009.
	C12	Yamagata, K., <i>et al.</i> , "Mutations In The Hepatocyte Nuclear Factor-1 $\alpha$ Gene In Maturity-Onset Diabetes Of The Young (MODY3)," <i>Nature</i> , 384: 455-458 (1996).
	C13	Froguel, Ph., <i>et al.</i> , "Close Linkage Of Glucokinase Locus On Chromosome 7p To Early-Onset Non-insulin-Dependent Diabetes Mellitus," <i>Nature</i> , 356: 162-164 (1992).
	C14	Antonetti, D. A., <i>et al.</i> , "Increased Expression of Mitochondrial-encoded Genes in Skeletal Muscle of Humans with Diabetes Mellitus," <i>J. Clinical Investigation</i> , 95: 1383-1388 (1995).
	C15	Little, M. H., <i>et al.</i> , "Dual Trafficking of Slit3 to Mitochondria and Cell Surface Demonstrates Novel Localization for Slit Protein," <i>Am. J. Physiol Cell Physiol</i> , 281: C486-C495 (2001).
	C16	Yuan, Shyng-Shiou, F., <i>et al.</i> , "Cloning and Functional Studies of a Novel Gene Aberrantly Expressed in RB-Deficient Embryos," <i>Developmental Biology</i> , 207: 62-75 (1999).
	C17	Maechler, P., and Wollheim, C. B., "Mitochondrial Function in Normal and Diabetic $\beta$ -cells," <i>Nature</i> , 414: 807-812 (2001).
	C18	Stein, E. and Tessier-Lavigne, M., "Hierarchical Organization of Guidance Receptors: Silencing of Netrin Attraction by Slit Through a Robo/DCC Receptor Complex," <i>Science</i> , 291: 1928-1938 (2001).

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	C19	Fricke, C., <i>et al.</i> , "Astray, A Zebrafish roundabout Homolog Required for Retinal Axon Guidance," <i>Science</i> , 292: 507-510 (2001).
	C20	Kang, J.S., <i>et al.</i> , "CDO, A Robo-related Cell Surface Protein That Mediates Myogenic Differentiation," <i>The Journal of Cell Biology</i> , 143(2): 403-413 (1998).
	C21	Shu, T. and Richards, L. J., "Crotical Axon Fuidance by the Glial Wedge During The Development of the Corpus Callosum," <i>The Journal of Neuroscience</i> , 21(8): 2749-2758 (2001).
	C22	Erschine, L., <i>et al.</i> , "Retinal Ganglion Cell Axon Guidance in the Mouse Optic Chiasm: Expression and Function of Robos and Slits," <i>The Journal of Neuroscience</i> , 20(13): 4975-4982 (2000).
	C23	Niclou, S. P., <i>et al.</i> , "Slit2 Is A Repellent for Retinal Ganglion Cell Axons," <i>The Journal of Neuroscience</i> , 20(13): 4962-4974 (2000).
	C24	Zou, Y., <i>et al.</i> , "Squeezing Axons Out of the Gray Matter: A Role for Slit and Semaphorin Proteins from Midline and Ventral Spinal Cord," <i>Cell</i> , 102: 363-375 (2000).
	C25	Brose, K., <i>et al.</i> , "Slit Proteins Bind Robo Receptors and Have an Evolutionary Conserved Role in Repulsive Axon Guidance," <i>Cell</i> , 96: 795-806 (1999).
	C26	Kidd, T., <i>et al.</i> , "Roundabout Controls Axon Crossing of the CNS Midline and Defines a Novel Subfamily of Evolutionary Conserved Guidance Receptors," <i>Cell</i> , 92: 205-215 (1998).
	C27	Kidd, T., <i>et al.</i> , "Slit Is The Midline Repellent For The Robo Receptor in <i>Drosophila</i> ," <i>Cell</i> , 96: 785-794 (1999).
	C28	Busfield, F., <i>et al.</i> , "A Genomewide Search for Type 2 Diabetes-Susceptibility Genes in Indigenous Australians," <i>Am. J. Hum. Genet.</i> , 70: 349-357 (2002).
	C29	Vionnet, N., <i>et al.</i> , "Genomewide Search for Type 2 Diabetes-Susceptibility Genes in French Whites: Evidence for a Novel Susceptibility Locus for Early-Onset Diabetes on Chromosome 3q27-qter and Independent Replication of a Type 2-Diabetes Locus on Chromosome 1q21-q24," <i>Am. J. Hum Genet.</i> , 67: 1470-1480 (2000).
	C30	Ge der Ehm, M., <i>et al.</i> , "Genomewide Search for Type 2 Diabetes Susceptibility Genes in Four American Populations," <i>Am. J. Hum. Genet.</i> , 66: 1871-1881 (2000).
	C31	Hanson, R. L., <i>et al.</i> , "An Autosomal Genomic Scan for Loci Linked to Type II Diabetes Mellitus and Body-Mass Index in Pima Indians," <i>Am. J. Hum. Genet.</i> , 63: 1130-1138 (1998).

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	C32	Duggirala, R., <i>et al.</i> , "A Major Locus for Fasting Insulin Concentrations and Insulin Resistance on Chromosome 6q with Strong Pleiotropic Effects on Obesity-Related Phenotypes in Nondiabetic Mexican Americans," <i>Am. J. Hum. Genet.</i> , 68: 1149-1164 (2001).
	C33	Baier, L., <i>et al.</i> , "A Calpain-10 Gene Polymorphism is Associated with Reduced Muscle mRNA levels and Insulin Resistance," <i>The Journal of Clinical Investigation</i> , 106: R69-R73 (2000).
	C34	Kristinsson, S. Y., <i>et al.</i> , "MODY in Iceland is Associated with Mutations in HNF-1a and a Novel Mutation in NeuroD1," <i>Diabetologia</i> , 44: 2098-2103 (2001).
	C35	Parker, A., <i>et al.</i> , "A Gene Conferring Susceptibility to Type 2 Diabetes in Conjunction With Obesity is Located on Chromosome 18p11," <i>Diabetes</i> , 50: 675-680 (2001).
	C36	Elbein, S., <i>et al.</i> , "A Genome-Wide Search for Type 2 Diabetes Susceptibility Genes in Utah Caucasians," <i>Diabetes</i> , 48: 1175-1182 (1999).
	C37	Sreenan, S. K., <i>et al.</i> , "Calpains Play a Role In Insulin Secretion and Action," <i>Diabetes</i> , 50: 2013-2020 (2001).
	C38	Lynn, S., <i>et al.</i> , "Variation in the Calpain-10 Gene Affects Blood Glucose Levels in the British Population," <i>Diabetes</i> , 51: 247-250 (2002).
	C39	Ronca, F., <i>et al.</i> , "Characterization of Slit Protein Interactions with Glypican-1," <i>The Journal of Biological Chemistry</i> , 276(31): 29141-29147 (2001).
	C40	Reynisdottir, I., <i>et al.</i> , "Localization of a Susceptibility Gene for Type 2 Diabetes to Chromosome 5q34-q35.2," <i>Am. J. Hum. Genet.</i> 73: 323-335 (2003).
	C41	Horikawa, J., <i>et al.</i> , "Mutation in Hepatocyte Nuclear Factor-1 $\beta$ gene (TCF2) associated with MODY," <i>Nature Genetics</i> , 17: 384-385 (1997).
	C42	Cox, J., <i>et al.</i> , "Loci on Chromosomes 2 (NIDDM1) and 15 Interact to Increase Susceptibility to Diabetes in Mexican Americans," <i>Nature Genetics</i> , 21: 213-215 (1999).
	C43	van den Ouweland, J.M.W., <i>et al.</i> , "Mutation in Mitochondrial tRNA <sup>Leu(UUR)</sup> Gene in a Large Pedigree with Maternity Transmitted Type II Diabetes Mellitus and Deafness," <i>Nature Genetics</i> , 1: 368-371 (1992).
	C44	Mahtani, M.M., <i>et al.</i> , "Mapping of a Gene for Type 2 Diabetes Associated with an Insulin Secretion Defect by a Genome Scan in Finnish Families," <i>Nature Genetics</i> , 14: 90-94 (1996).

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	C45	Hanis, C.L., <i>et al.</i> , "A Genome-wide Search for Human Non-Insulin-Dependent (Type 2) Diabetes Genes Reveals a Major Susceptibility Locus on Chromosome 2," <i>Nature Genetics</i> , 13: 161-166 (1996).
	C46	Horikawa, Y., <i>et al.</i> , "Genetic Variation In The Gene Encoding Calpain-10 is Associated with Type 2 Diabetes Mellitus," <i>Nature Genetics</i> , 26: 163-175 (2000).
	C47	Marillat, V., <i>et al.</i> , "Spatiotemporal Expression Patterns of <i>slit</i> and <i>robo</i> Genes in the Rat Brain," <i>The Journal of Comparative Neurology</i> , 442: 130-155 (2002).
	C48	Lee, H.K., <i>et al.</i> , "Decreased Mitochondrial DNA Content in Peripheral Blood Precedes The Development of Non-Insulin-Dependent Diabetes Mellitus," <i>Diabetes Research and Clinical Practice</i> , 42: 161-167 (1998).
	C49	Nakayama, M., <i>et al.</i> , "Identification of High-Molecular-Weight Proteins with Multiple EGF-like Motifs by Motif-Trap Screening," <i>Genomics</i> , 51: 27-34 (1998).
	C50	Yoshioka, M., <i>et al.</i> , "A Novel Locus, <i>Mody4</i> , Distal to D7Mit189 on Chromosome 7 Determines Early-Onset NIDDM in Nonobese C57BL/6 (Akita) Mutant Mice," <i>Diabetes</i> , 46(5): 887-894 (1997).

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